



The present invention relates to a novel tissue culture system that provides

for the long term culture of proliferating hepatocytes that retain hepatic function/ coll author

Disclosed are methods and compositions for ex vivo-culturing of nepatocytes and nonparenchymal cells on a matrix coated with a molecule that promotes cell adhesion,

proliferation or survival, in the presence of growth factors, resulting in a long-term 5 culture of proliferating hepatocytes that retain hepatic function. The co-culturing method results in the formation of matrix/hepatic cell clusters that may be mixed with a second structured or scaffold matrix that provides a three-dimensional structural support to form 6 structures analogous to liver tissue counterparts. The hepatic cell culture system can be

10 used to form bio-artificial livers through which a subjects blood is perfused. Alternatively, the novel hepatic cell culture system may be implanted into the body of a recipient host having a hepatic disorder. Such hepatic disorders, include, for example,

cirrhosis of the liver, induced hepatitis, chronic hepatitis, primary sclerosing cholangitis

and alpha<sub>1</sub> antitrypsin deficiency.